FORM PTO-1390 (Modified) U.S. DEP: STMENT OF COMM. PATENT AND TRADEMARK OFFICE (REV. 11-2000)

TRANSMITTAL LETTER TO THE UNITED STATES DESIGNATED/ELECTED OFFICE (DO/EO/US)

CONCERNING A FILING UNDER 35 U.S.C. 371

17703 (PC27210A)

U.S. APPLICATION NO. (IF KNOWN, SEE 37 CFR 10/500606

INTERNATIONAL APPLICATION NO. PCT/EP02/13092

INTERNATIONAL FILING DATE

18 December 2002 (18.12.2002)

PRIORITY DATE CLAIMED 2 January 2002 (02.01.2002)

TITLE OF INVENTION

COMBINED THERAPY AGAINST TUMORS COMPRISING SUBSTITUTED ACRYLOYL DISTAMYCIN DERIVATIVES AND PROTEIN KINASE (SERINE/THREONINE KINASE) INHIBITORS

APPLICANT(S) FOR DO/EO/US

Maria Geroni, Camilla Fowst and Paolo Cozzi

| | | · | | | | |
|-------|-------------|---|--|--|--|--|
| Appli | icant h | nerewith submits to the United States Designated/Elected Office (DO/EO/US) the following items and other information: | | | | |
| 1. | × | This is a FIRST submission of items concerning a filing under 35 U.S.C. 371. | | | | |
| 2. | | This is a SECOND or SUBSEQUENT submission of items concerning a filing under 35 U.S.C. 371. | | | | |
| 3. | | This is an express request to begin national examination procedures (35 U.S.C. 371(f)). The submission must include items (5), (6), (9) and (24) indicated below. | | | | |
| 4. | \boxtimes | The US has been elected by the expiration of 19 months from the priority date (Article 31). | | | | |
| 5. | \boxtimes | A copy of the International Application as filed (35 U.S.C. 371 (c) (2)) | | | | |
| | | a. is attached hereto (required only if not communicated by the International Bureau). | | | | |
| | | b. 🛛 has been communicated by the International Bureau. | | | | |
| | | c. \square is not required, as the application was filed in the United States Receiving Office (RO/US). | | | | |
| 6. | | An English language translation of the International Application as filed (35 U.S.C. 371(c)(2)). | | | | |
| | | a. is attached hereto. | | | | |
| | | b. has been previously submitted under 35 U.S.C. 154(d)(4). | | | | |
| 7. | \boxtimes | Amendments to the claims of the International Application under PCT Article 19 (35 U.S.C. 371 (c)(3)) | | | | |
| | | a. are attached hereto (required only if not communicated by the International Bureau). | | | | |
| | | b. have been communicated by the International Bureau. | | | | |
| | | c. \square have not been made; however, the time limit for making such amendments has NOT expired. | | | | |
| | | d. 🛮 have not been made and will not be made. | | | | |
| 8. | | An English language translation of the amendments to the claims under PCT Article 19 (35 U.S.C. 371(c)(3)). | | | | |
| 9. | | An oath or declaration of the inventor(s) (35 U.S.C. 371 (c)(4)). | | | | |
| 10. | | An English language translation of the annexes to the International Preliminary Examination Report under PCT Article 36 (35 U.S.C. 371 (c)(5)). | | | | |
| 11. | \boxtimes | A copy of the International Preliminary Examination Report (PCT/IPEA/409). | | | | |
| 12. | \boxtimes | A copy of the International Search Report (PCT/ISA/210). | | | | |
| It | ems 1 | 3 to 20 below concern document(s) or information included: | | | | |
| 13. | | An Information Disclosure Statement under 37 CFR 1.97 and 1.98. | | | | |
| 14. | | An assignment document for recording. A separate cover sheet in compliance with 37 CFR 3.28 and 3.31 is included. | | | | |
| 15. | | A FIRST preliminary amendment. | | | | |
| 16. | | A SECOND or SUBSEQUENT preliminary amendment. | | | | |
| 17. | | A substitute specification. | | | | |
| 18. | | A change of power of attorney and/or address letter. | | | | |
| 19. | | A computer-readable form of the sequence listing in accordance with PCT Rule 13ter.2 and 35 U.S.C. 1.821 - 1.825. | | | | |
| 20. | | A second copy of the published international application under 35 U.S.C. 154(d)(4). | | | | |
| 21. | | A second copy of the English language translation of the international application under 35 U.S.C. 154(d)(4). | | | | |
| 22. | \boxtimes | Express Mail Label No. <u>EV213901318US</u> | | | | |
| 23. | \boxtimes | Other items or information: | | | | |
| | | Courtesy copy of International Application Assignee: PHARMACIA ITALIA SPA of Milan, Italy | | | | |

TTORNEY'S DOCKET NUMBER JUN 2004

| | 10/500606 PCT/EP02/13092 | | | 17703 (PC27210A) | | | |
|--|--|---------------------|----------------|------------------------|-------------|--|--|
| 24. The following fees are submitted:. | | CALCULATION | S PTO USE ONLY | | | | |
| BASIC NATIONAL FEE (37 CFR 1.492 (a) (1) - (5)): Neither international preliminary examination fee (37 international search fee (37 CFR 1.445(a)(2)) paid to and International Search Report not prepared by the E | 80.00 | | | | | | |
| ☑ International preliminary examination fee (37 CFR 1.4 USPTO but International Search Report prepared by t | 20.00 | | | | | | |
| ☐ International preliminary examination fee (37 CFR 1.4 but international search fee (37 CFR 1.445(a)(2)) paid | 70.00 | | | | | | |
| ☐ International preliminary examination fee (37 CFR 1.4 but all claims did not satisfy provisions of PCT Article | 30.00 | | | | | | |
| ☐ International preliminary examination fee (37 CFR 1.4 and all claims satisfied provisions of PCT Article 33(1) | 00.00 | | | | | | |
| ENTER APPROPRIATE I | | \$920.00 | | | | | |
| Surcharge of \$130.00 for furnishing the oath or declaration la months from the earliest claimed priority date (37 CFR 1.492 | ter than |) 🛛 | 30 | \$130.00 | | | |
| | UMBER EXTRA | RA RATE | | | | | |
| Total claims 27 - 20 = | 7 | x \$18. | 00 | \$126.00 | | | |
| Independent claims 6 - 3 = | 3 | x \$86. | 00 | \$258.00 | | | |
| Multiple Dependent Claims (check if applicable). | | Ø | | \$290.00 | | | |
| TOTAL OF ABO | | | = | \$1,724.00 | | | |
| Applicant claims small entity status. See 37 CFR 1.27. Treduced by 1/2. | The fees indicated above | are | | \$0.00 | | | |
| | SUB | ΓΟΤΑL | , = | \$1,724.00 | | | |
| Processing fee of \$130.00 for furnishing the English translation months from the earliest claimed priority date (37 CFR 1.492) | on later than \Box 20.2. (f)). |) 🗆 : | 30 + | \$0.00 | | | |
| TO | TAL NATIONAL | FEE | = | \$1,724.00 | | | |
| Fee for recording the enclosed assignment (37 CFR 1.21(h)). accompanied by an appropriate cover sheet (37 CFR 3.28, 3.3). | The assignment must b | e). | | \$0.00 | | | |
| TOTAL FEES ENCLOSED = | | | | \$1,724.00 | | | |
| | | | | Amount to be: refunded | \$ | | |
| | | | | charged | \$ | | |
| a. A check in the amount of \$1,724.00 to cover the above fees is enclosed. b. Please charge my Deposit Account No. in the amount of to cover the above fees. | | | | | | | |
| c. The Director is hereby authorized to charge an to Deposit Account No. 19-1013/SSMP | | | | | | | |
| d. Fees are to be charged to a credit card. WARNING: Information on this form may become public. Credit card information should not be included on this form. Provide credit card information and authorization on PTO-2038. | | | | | | | |
| NOTE: Where an appropriate time limit under 37 CFR 1 1.137(a) or (b)) must be filed and granted to restore the ap | .494 or 1.495 has not b plication to pending st | een met, a | petitio | n to revive (37 CFI | | | |
| SEND ALL CORRESPONDENCE TO: | | | | | | | |
| Peter I. Bernstein, Esq. | | SIGNAT | TURE | | | | |
| Registration No. 43,497 SCULLY, SCOTT, MURPHY & PRESSER | | | | | | | |
| 400 Garden City Plaza | | Peter I. Bernstein | | | | | |
| Garden City, New York 11530 | } | NAME | | | | | |
| (516) 742-4343 | | 43,497 | | | | | |
| | | REGISTRATION NUMBER | | | | | |
| | | June 30, 2004 | | | | | |
| | | DATE | | | | | |

10/500606

| | RTIFICATE OF I | Rec'd PCT/PPocks No JUN 2004 17703 (PC27210A) | | | | | | | | | |
|-----------------------|---|--|---|---|--|--|--|--|--|--|--|
| Serial No. Unassigned | | Filing Date June 30, 2004 | Examiner . Unassigned | Group Art Unit Unassigned | | | | | | | |
| Inver | Invention: COMBINED THERAPY AGAINST TUMORS COMPRISING SUBSTITUTED ACRYLOYL DISTAMYCIN DERIVATIVES AND PROTEIN KINASE (SERINE/THREONINE KINASE) INHIBITORS | | | | | | | | | | |
| | I hereby certify that the following correspondence: New U.S. Application Under 35 USC 371 | | | | | | | | | | |
| | (Identify type of correspondence) is being deposited with the United States Postal Service "Express Mail Post Office to Addressee" service under 37 CFR 1.10 in an envelope addressed to: Commissioner for Patents, P.O. Box 1450, Alexandria, VA 22313-1450 on | | | | | | | | | | |
| | | | Peter I. Ben (Typed or Printed Name of Person (Signature of Person Mailin EV2139013 ("Express Mail" Mailing | Mailing Correspondence) Ing Correspondence) 318US | | | | | | | |

Note: Each paper must have its own certificate of mailing.